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Milan, Dec 12, 2018

Phillip Steindel, Ph.D.

Review Editor, Journal of Visualized Experiments (JoVE)

Cc:

Ronald Myers, PhD.

Senior Science Editor, Journal of Visualized Experiments (JoVE)

1 Alewife Center, Suite 200, Cambridge, MA 02140

RE: Resubmission of JoVE58507 ENDOTOXIN ACTIVITY ASSAY FOR THE
DETECTION OF WHOLE BLOOD ENDOTOXEMIA IN CRITICALLY ILL PATIENTS,

Response to Review - Cover Letter

Dear Dr. Steindel,

We are pleased to resubmit for publication a new version of our manuscript, modified according to the Editor's recommendations. Particularly, point-by-point:

1. You have indicated in your ALA that you want Open Access but checked Standard in Editorial Manager-which are you choosing? If standard, please sign a new ALA (attached) and include it with your submission.

A new ALA has been attached to the resubmission. We confirm we want Standard Access.

2. EAA is a commercial term (see <http://www.spectraldx.com/ea.html>, e.g.); please use another abbreviation for this assay.

Throughout the revised version of the manuscript, we referred to the measurement of "Endotoxin Activity (EA)". No brand name has been used.

3. Please include an ethics statement before the numbered protocol steps, indicating that the protocol follows the guidelines of your institution's human research ethics committee.

Added. See Page 3, lines 102-105.

4.

4. 1: This section should be in the imperative.

The section has been modified as requested (Page 3, from line 110)

5. 2.10: How long do you vortex here?

10 seconds. The text has been changed accordingly.

6. 2.10.2: What does 'for duplicate tubes 1, 2 and 3' mean-repeat the readings for those tubes?

Each EA assay is run in duplicates, meaning the same blood sample is tested through chemiluminometry in two separate sets of tubes (tube #1, tube #2, tube #3 AND tube#1-duplicate, tube#2-duplicate, tube#3-duplicate). The two resulting EAs are averaged. This is repeatedly remarked in the protocol description (1.3, 2.2, 2.5, 2.6, 2.7, 2.10.2, 2.11).

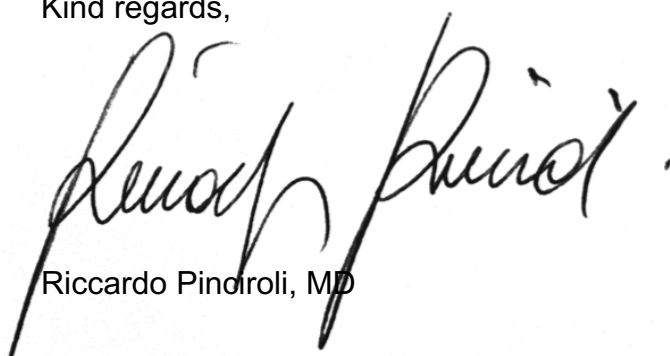
7. 3: This section probably belongs (in paragraph form) in the results section; otherwise it should be in the imperative and entirely numbered steps.

We thank the editor for the useful suggestion. After proper rephrasing, the paragraph was moved to the Representative Results section.

8. Figure 1: It is unclear why this is here; the video will probably convey all essential information about the instrument.

The figure was added during the review process following reviewer #1/comment #3. However, we agree with the editor and removed the figure representing the instrument.

Kind regards,



Riccardo Pinioli, MD